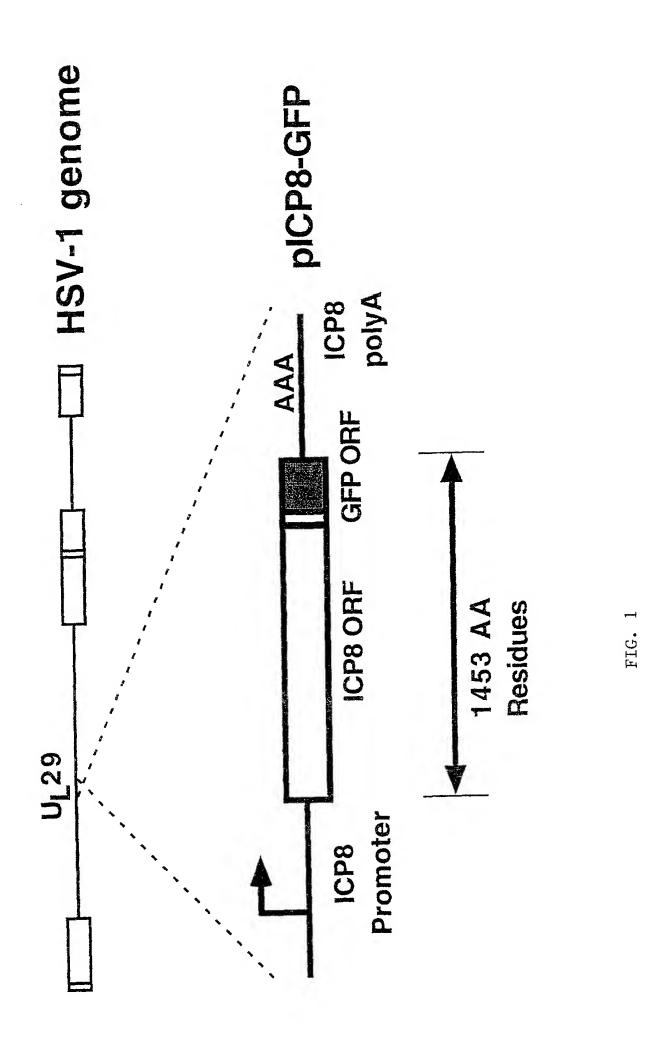
Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, et al.



App No.:	0725.1059-001
Title: "Repli	cation-Competent Virus"
Inventors:	David M. Knipe, et al.
	-

 8.5×10^8

 8.8×10^8

 7.7×10^8

 6.6×10^7

 4.6×10^{5}

20

ICP8-GFP

KOS 1.1

20

KOS 1.1

 6.8×10^8

 1.5×10^8

 9.8×10^4

ICP8-GFP

S2

YIELD (PFU//ML)^a

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Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, et al.

FIG. 3A

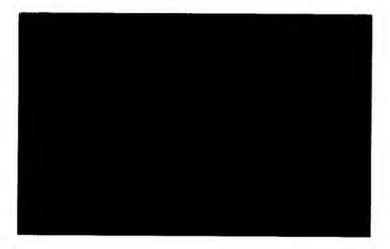


FIG. 3B

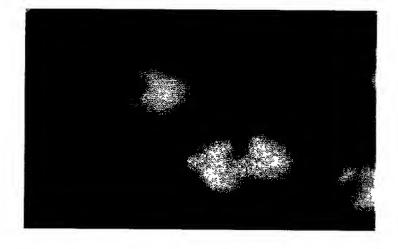


FIG. 3C

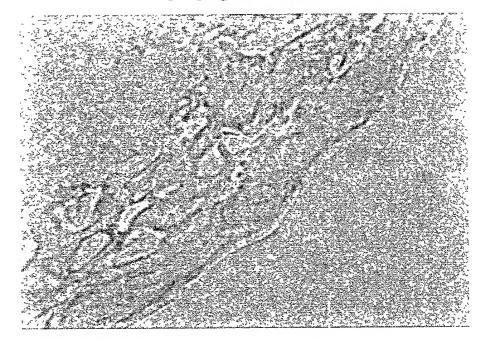


App No.: 0725.1059-001
Title: "Replication-Competent Virus ..."
Inventors: David M. Knipe, et al.

FIG. 4A



FIG. 4B



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Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, *et al.*

atg gag aca aag ccc aag acg gca acc acc atc aag gtc ccc ccc ggg 48 ccc ctg gga tac gtg tac gct cgc gcg tgt ccg tcc gaa ggc atc gag 96 ctt ctg gcg tta ctg tcg gcg cgc agc ggc gat gcc gac gtc gcc gtg 144 gcg ccc ctg gtc gtg ggc ctg acc gtg gag agc ggc ttt gag gcc aac 192 gta gcc gtg gtc gtg ggt tct cgc acg acg ggg ctc ggg ggt acc gcg 240 gtg tcc ctg aaa ctg acg cca tcg cac tac agc tcg tcc gtg tac gtc 288 ttt cac ggc ggc cgg cac ctg gac ccc agc acc cag gcc cca aac ctg 336 acg cga ctc tgc gag cgg gca cgc cgc cat ttt ggc ttt tcg gac tac 384 acc ccc cgg ccc ggc gac ctc aaa cac gag acg acg ggg gag gcg ctg 432 tgt gag cgc ctc ggc ctg gac ccg gac cgc gcc ctc ctg tat ctg gtc 480 gtt acc gag ggc ttc aag gag gcc gtg tgc atc aac acc ttt ctg 528 cac ctg gga ggc tcg gac aag gta acc ata ggc ggg gcg gag gtg cac 576 cgc ata ccc gtg tat ccg ttg cag ctg ttc atg ccg gat ttt agc cgg 624 gtc atc gcc gag ccg ttc aac gcc aac cac cga tcg atc ggg gag aat 672 ttt acc tac ccg ctt ccg ttt ttt aac cgc ccc ctc aac cgc ctc ctg 720 ttc gag gcg gtc gtg gga ccc gcc gcc gtg gca ctg cga tgc cga aac 768 gtg gac gcc gtg gcc cgc gcc gcc cac ctg gcg ttt gac gaa aac 816 cac gag ggc gcc gcc ccc gcc gac att acg ttc acg gcc ttc gaa 864 gcc agc cag ggt aag acc ccg cgg ggt ggg cgc gac ggc ggc aag 912 ggc ccg gcg ggc ggg ttc gaa cag cgc ctg gcc tcc gtc atg gcc gga 960 gac gcc gcc ctg gcc ctc gag tct atc gtg tcg atg gcc gtc ttc gac 1008 gag ccg ccc acc gac atc tcc gcg tgg ccg ctg tgc gag ggc cag gac 1056 acg gcc gcg gcc cgc gcc aac gcc gtc ggg gcg tac ctg gcg cgc gcc 1104 gcg gga ctc gtg ggg gcc atg gta ttt agc acc aac tcg gcc ctc cat 1152 ctc acc gag gtg gac gac gcc ggt ccg gcg gac cca aag gac cac agc 1200 aaa ccc tcc ttt tac cgc ttc ttc ctc gtg ccc ggg acc cac gtg gcg 1248 gcc aac cca cag gtg gac cgc gag gga cac gtg gtg ccc ggg ttc gag 1296 ggt cgg ccc acc gcg ccc ctc gtc ggc gga acc cag gaa ttt gcc ggc 1344 gag cac ctg gcc atg ctg tgt ggg ttt tcc ccg gcg ctg ctg gcc aag 1392 atg ctg ttt tac ctg gag cgc tgc gac ggc gtg atc gtc ggg cgc 1440 FIG. 6A

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Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, et al.

cag gag atg gac gtg ttt cga tac gtc gcg gac tcc aac cag acc gac 1488 gtg ccc tgc aac ctg tgc acc ttc gac acg cgc cac gcc tgc gta cac 1536 acg acg ctc atg cgc ctc cgg gcg cgc cat ccc aag ttc gcc agc gcc 1584 gcc cgc gga gcc atc ggc gtc ttc ggg acc atg aac agc atg tac agc 1632 gac tgc gac gtg ctg gga aac tac gcc gcc ttc tcg gcc ctg aag cgc 1680 gcg gac gga tcc gag acc gcc cgg acc atc atg cag gag acg tac cgc 1728 gcg gcg acc gag cgc gtc atg gcc gaa ctc gag acc ctg cag tac gtg 1776 gac dag gcg gtc ccc acg gcc atg ggg cgg ctg gag acc atc acc 1824 aac cgc gag gcc ctg cat acg gtg gtg aac aac gtc agg cag gtc gtg 1872 gac cgc gag gtg gag cag ctg atg cgc aac ctg gtg gag ggg agg aac 1920 ttc aag ttt cgc gac ggt ctg ggc gag gcc aac cac gcc atg tcc ctg 1968 acg ctg gac ccg tac gcg tgc ggg cca tgc ccc ctg ctt cag ctt ctc 2016 ggg cgg cga tcc aac ctc gcc gtg tat cag gac ctg gcc ctg agc cag 2064 tgc cac ggg gtg ttc gcc ggg cag tcg gtc gag ggg cgc aac ttt cgc 2112 aat caa ttc caa ccg gtg ctg cgg cgc gtg atg gac atg ttt aac 2160 aac ggg ttt ctg tcg gcc aaa acg ctg acg gtc gcg ctc tcg gag ggg 2208 geg get ate tge gee eec age eta aeg gee gge eag aeg gee eec gee 2256 gag age age tte gag gge gae gtt gee ege gtg ace etg ggg ttt eee 2304 aag gag ctg cgc gtc aag agc cgc gtg ttg ttc gcg ggc gcg agc gcc 2352 aac gcg tcc gag gcc gcc aag gcg cgg gtc gcc agc ctc cag agc gcc 2400 tac cag aag ccc gac aag cgc gtg gac atc ctc ctc gga ccg ctg ggc 2448 ttt ctg ctg aag cag ttc cac gcg gcc atc ttc ccc aac ggc aag ccc 2496 ccg ggg tcc aac cag ccg aac ccg cag tgg ttc tgg acg gcc ctc caa 2544 cgc aac cag ctt ccc gcc cgg ctc ctg tcg cgc gag gac atc gag acc 2592 atc gcg ttc att aaa aag ttt tcc ctg gac tac ggc gcg ata aac ttt 2640 att aac ctg gcc ccc aac aac gtg agc gag ctg gcg atg tac tac atg 2688 gca aac cag att ctg cgg tac tgc gat cac tcg aca tac ttc atc aac 2736 acc ctc acg gcc atc acc gcg ggg tcc cgc cgt ccc ccc agc gtg cag 2784 geg geg gee geg tgg tee geg cag gge ggg geg gge etg gag gee ggg 2832 gcc cgc gcg ctg atg gac gcc gtg gac gcg cat ccg ggc gcg tgg acg 2880 tee atg tte gee age tge aac etg etg egg eee gte atg geg ege 2928

Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, et al.

ccc atg gtc gtg ttg ggg ttg agc atc agc aaa tac tac ggc atg gcc 2976 3024 ggc aac gac cgt gtg ttt cag gcc ggg aac tgg gcc agc ctg atg ggc qgc aaa aac gcg tgc ccg ctc ctt att ttt gac cgc acc cgc aag ttc 3072 gtc ctg gcc tgt ccc cgg gcc ggg ttt gtg tgc gcg gcc tcg aac ctc 3120 ggc ggc gga gcg cac gaa agc tcg ctg tgc gag cag ctc cgg ggc att 3168 atc tec gag gge ggg gee gte gee agt age gtg tte gtg geg ace 3216 3264 gtg aaa agc ctg ggg ccc cgc acc cag cag ctg cag atc gag gac tgg ctg gcg ctc ctg gag gac gag tac cta agc gag gag atg atg gag ctg 3312 acc gcg cgt gcc ctg gag cgc ggc aac ggc gag tgg tcg acg gac gcg 3360 3408 gee etg gag gtg geg cae gag gee gag gee eta gte age caa ete gge aac gcc ggg gag gtg ttt aac ttt ggg gat ttt ggc tgc gag gac gac 3456 aac gcg acg ccg ttc ggc ggc ccg ggg gcc ccg gga ccg gca ttt gcc 3504 ggc cgc aaa cgg gcg ttc cac ggg gat gac ccg ttt ggg gag ggg ccc 3552 3600 ccc gac aaa aag gga gac ctg acg ttg gat atg ctg aga ggg gtt ggg 3648 ggg tgg ggg aac cta gag tcg acc cgg gcg gcc gcc gcc acc atg agc aag ggc gag gaa ctg ttc act ggc gtg gtc cca att ctc gtg gaa ctg 3696 3744 gat ggc gat gtg aat ggg cac aaa ttt tct gtc agc gga gag ggt gaa ggt gat gcc aca tac gga aag ctc acc ctg aaa ttc atc tgc acc act 3792 gga aag ctc cct gtg cca tgg cca aca ctg gtc act acc ttc acc tat 3840 ggc gtg cag tgc ttt tcc aga tac cca gac cat atg aag cag cat gac 3888 3936 ttt ttc aag agc gcc atg ccc gag ggc tat gtg cag gag aga acc atc ttt ttc aaa gat gac ggg aac tac aag acc cgc gct gaa gtc aag ttc 3984 4032 gaa ggt gac acc ctg gtg aat aga atc gag ttg aag ggc att gac ttt aag gaa gat gga aac att ctc ggc cac aag ctg gaa tac aac tat aac 4080 tcc cac aat gtg tac atc atg gcc gac aag caa aag aat ggc atc aag 4128 gtc aac ttc aag atc aga cac aac att gag gat gga tcc gtg cag ctg 4176 gcc gac cat tat caa cag aac act cca atc ggc gac ggc cct gtg ctc 4224 ctc cca gac aac cat tac ctg tcc acc cag tct gcc ctg tct aaa gat 4272 ccc aac gaa aag aga gac cac atg gtc ctg ctg gag ttt gtg acc gct 4320 gct ggg atc aca cat ggc atg gac gag ctg tac aag tga 4359 FIG. 6C

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Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, *et al*.

Met Glu Thr Lys Pro Lys Thr Ala Thr Thr Ile Lys Val Pro Pro Gly Pro Leu Gly Tyr Val Tyr Ala Arg Ala Cys Pro Ser Glu Gly Ile Glu 25 Leu Leu Ala Leu Leu Ser Ala Arg Ser Gly Asp Ala Asp Val Ala Val Ala Pro Leu Val Val Gly Leu Thr Val Glu Ser Gly Phe Glu Ala Asn Val Ala Val Val Gly Ser Arg Thr Thr Gly Leu Gly Gly Thr Ala 70 75 Val Ser Leu Lys Leu Thr Pro Ser His Tyr Ser Ser Ser Val Tyr Val 90 Phe His Gly Gly Arg His Leu Asp Pro Ser Thr Gln Ala Pro Asn Leu 100 105 Thr Arg Leu Cys Glu Arg Ala Arg Arg His Phe Gly Phe Ser Asp Tyr 115 120 Thr Pro Arg Pro Gly Asp Leu Lys His Glu Thr Thr Gly Glu Ala Leu 135 140 Cys Glu Arg Leu Gly Leu Asp Pro Asp Arg Ala Leu Leu Tyr Leu Val 150 155 Val Thr Glu Gly Phe Lys Glu Ala Val Cys Ile Asn Asn Thr Phe Leu His Leu Gly Gly Ser Asp Lys Val Thr Ile Gly Gly Ala Glu Val His 180 185 Arg Ile Pro Val Tyr Pro Leu Gln Leu Phe Met Pro Asp Phe Ser Arg 200 205 Val Ile Ala Glu Pro Phe Asn Ala Asn His Arg Ser Ile Gly Glu Asn 220 215 Phe Thr Tyr Pro Leu Pro Phe Phe Asn Arg Pro Leu Asn Arg Leu Leu 230 235 Phe Glu Ala Val Val Gly Pro Ala Ala Val Ala Leu Arg Cys Arg Asn 245 250 Val Asp Ala Val Ala Arg Ala Ala Ala His Leu Ala Phe Asp Glu Asn 260 265 270 His Glu Gly Ala Ala Leu Pro Ala Asp Ile Thr Phe Thr Ala Phe Glu 280 Ala Ser Gln Gly Lys Thr Pro Arg Gly Gly Arg Asp Gly Gly Lys 295 Gly Pro Ala Gly Gly Phe Glu Gln Arg Leu Ala Ser Val Met Ala Gly 310 315 Asp Ala Ala Leu Ala Leu Glu Ser Ile Val Ser Met Ala Val Phe Asp 330 Glu Pro Pro Thr Asp Ile Ser Ala Trp Pro Leu Cys Glu Gly Gln Asp 340 345 350 Thr Ala Ala Arg Ala Asn Ala Val Gly Ala Tyr Leu Ala Arg Ala 360 Ala Gly Leu Val Gly Ala Met Val Phe Ser Thr Asn Ser Ala Leu His 375 Leu Thr Glu Val Asp Asp Ala Gly Pro Ala Asp Pro Lys Asp His Ser 385 390 395 400 Lys Pro Ser Phe Tyr Arg Phe Phe Leu Val Pro Gly Thr His Val Ala 410

Title: "Replication-Competent Virus ..." Inventors: David M. Knipe, et al.

Ala Asn Pro Gln Val Asp Arg Glu Gly His Val Val Pro Gly Phe Glu 420 425 Gly Arg Pro Thr Ala Pro Leu Val Gly Gly Thr Gln Glu Phe Ala Gly 440 Glu His Leu Ala Met Leu Cys Gly Phe Ser Pro Ala Leu Leu Ala Lys 455 Met Leu Phe Tyr Leu Glu Arg Cys Asp Gly Gly Val Ile Val Gly Arg 470 475 Gln Glu Met Asp Val Phe Arg Tyr Val Ala Asp Ser Asn Gln Thr Asp 485 Val Pro Cys Asn Leu Cys Thr Phe Asp Thr Arg His Ala Cys Val His 505 Thr Thr Leu Met Arg Leu Arg Ala Arg His Pro Lys Phe Ala Ser Ala 515 520 Ala Arg Gly Ala Ile Gly Val Phe Gly Thr Met Asn Ser Met Tyr Ser 535 Asp Cys Asp Val Leu Gly Asn Tyr Ala Ala Phe Ser Ala Leu Lys Arg Ala Asp Gly Ser Glu Thr Ala Arg Thr Ile Met Gln Glu Thr Tyr Arg 565 570 Ala Ala Thr Glu Arg Val Met Ala Glu Leu Glu Thr Leu Gln Tyr Val 585 Asp Gln Ala Val Pro Thr Ala Met Gly Arg Leu Glu Thr Ile Ile Thr 600 605 Asn Arg Glu Ala Leu His Thr Val Val Asn Asn Val Arg Gln Val Val 615 Asp Arg Glu Val Glu Gln Leu Met Arg Asn Leu Val Glu Gly Arg Asn 630 635 Phe Lys Phe Arg Asp Gly Leu Gly Glu Ala Asn His Ala Met Ser Leu 645 650 Thr Leu Asp Pro Tyr Ala Cys Gly Pro Cys Pro Leu Leu Gln Leu Leu 660 665 Gly Arg Arg Ser Asn Leu Ala Val Tyr Gln Asp Leu Ala Leu Ser Gln 680 Cys His Gly Val Phe Ala Gly Gln Ser Val Glu Gly Arg Asn Phe Arg 695 700 Asn Gln Phe Gln Pro Val Leu Arg Arg Val Met Asp Met Phe Asn 710 715 Asn Gly Phe Leu Ser Ala Lys Thr Leu Thr Val Ala Leu Ser Glu Gly 725 730 Ala Ala Ile Cys Ala Pro Ser Leu Thr Ala Gly Gln Thr Ala Pro Ala 740 745 Glu Ser Ser Phe Glu Gly Asp Val Ala Arg Val Thr Leu Gly Phe Pro 760 Lys Glu Leu Arg Val Lys Ser Arg Val Leu Phe Ala Gly Ala Ser Ala 770 775 Asn Ala Ser Glu Ala Ala Lys Ala Arg Val Ala Ser Leu Gln Ser Ala 790 795 Tyr Gln Lys Pro Asp Lys Arg Val Asp Ile Leu Leu Gly Pro Leu Gly 805 810 Phe Leu Leu Lys Gln Phe His Ala Ala Ile Phe Pro Asn Gly Lys Pro 825 Pro Gly Ser Asn Gln Pro Asn Pro Gln Trp Phe Trp Thr Ala Leu Gln 835 840 Arg Asn Gln Leu Pro Ala Arg Leu Leu Ser Arg Glu Asp Ile Glu Thr 855

FIG. 7B

App No.: 0725.1059-001

Title: "Replication-Competent Virus ..."

Inventors: David M. Knipe, et al.

Ile Ala Phe Ile Lys Lys Phe Ser Leu Asp Tyr Gly Ala Ile Asn Phe Ile Asn Leu Ala Pro Asn Asn Val Ser Glu Leu Ala Met Tyr Tyr Met Ala Asn Gln Ile Leu Arg Tyr Cys Asp His Ser Thr Tyr Phe Ile Asn Thr Leu Thr Ala Ile Ile Ala Gly Ser Arg Arg Pro Pro Ser Val Gln Ala Ala Ala Trp Ser Ala Gln Gly Gly Ala Gly Leu Glu Ala Gly Ala Arg Ala Leu Met Asp Ala Val Asp Ala His Pro Gly Ala Trp Thr Ser Met Phe Ala Ser Cys Asn Leu Leu Arg Pro Val Met Ala Ala Arg Pro Met Val Val Leu Gly Leu Ser Ile Ser Lys Tyr Tyr Gly Met Ala Gly Asn Asp Arg Val Phe Gln Ala Gly Asn Trp Ala Ser Leu Met Gly Gly Lys Asn Ala Cys Pro Leu Leu Ile Phe Asp Arg Thr Arg Lys Phe Val Leu Ala Cys Pro Arg Ala Gly Phe Val Cys Ala Ala Ser Asn Leu Gly Gly Gly Ala His Glu Ser Ser Leu Cys Glu Gln Leu Arg Gly Ile Ile Ser Glu Gly Gly Ala Ala Val Ala Ser Ser Val Phe Val Ala Thr Val Lys Ser Leu Gly Pro Arg Thr Gln Gln Leu Gln Ile Glu Asp Trp Leu Ala Leu Leu Glu Asp Glu Tyr Leu Ser Glu Glu Met Met Glu Leu Thr Ala Arg Ala Leu Glu Arg Gly Asn Gly Glu Trp Ser Thr Asp Ala Ala Leu Glu Val Ala His Glu Ala Glu Ala Leu Val Ser Gln Leu Gly Asn Ala Gly Glu Val Phe Asn Phe Gly Asp Phe Gly Cys Glu Asp Asp Asn Ala Thr Pro Phe Gly Gly Pro Gly Ala Pro Gly Pro Ala Phe Ala Gly Arg Lys Arg Ala Phe His Gly Asp Pro Phe Gly Glu Gly Pro Pro Asp Lys Cly Asp Leu Thr Leu Asp Met Leu Arg Gly Val Gly Gly Trp Gly Asn Leu Glu Ser Thr Arg Ala Ala Ala Thr Met Ser Lys Gly Glu Leu Phe Thr Gly Val Val Pro Ile Leu Val Glu Leu Asp Gly Asp Val Asn Gly His Lys Phe Ser Val Ser Gly Glu Gly Glu Gly Asp Ala Thr Tyr Gly Lys Leu Thr Leu Lys Phe Ile Cys Thr Thr Gly Lys Leu Pro Val Pro Trp Pro Thr Leu Val Thr Thr Phe Thr Tyr Gly Val Gln Cys Phe Ser Arg Tyr Pro Asp His Met Lys Gln His Asp Phe Phe Lys Ser Ala Met Pro Glu Gly Tyr Val Gln Glu Arg Thr Ile Phe Phe Lys Asp Asp Gly Asn Tyr Lys Thr Arg Ala Glu Val Lys Phe

App No.: 0725.1059-001

Title: "Replication-Competent Virus ..."

Inventors: David M. Knipe, *et al*.

Glu Gly Asp Thr Leu Val Asn Arg Ile Glu Leu Lys Gly Ile Asp Phe 1335 Lys Glu Asp Gly Asn Ile Leu Gly His Lys Leu Glu Tyr Asn Tyr Asn 1350 1355 1360 Ser His Asn Val Tyr Ile Met Ala Asp Lys Gln Lys Asn Gly Ile Lys 1365 1370 Val Asn Phe Lys Ile Arg His Asn Ile Glu Asp Gly Ser Val Gln Leu 1380 1385 Ala Asp His Tyr Gln Gln Asn Thr Pro Ile Gly Asp Gly Pro Val Leu 1395 1400 Leu Pro Asp Asn His Tyr Leu Ser Thr Gln Ser Ala Leu Ser Lys Asp 1410 1415 Pro Asn Glu Lys Arg Asp His Met Val Leu Leu Glu Phe Val Thr Ala 1425 1430 1435 Ala Gly Ile Thr His Gly Met Asp Glu Leu Tyr Lys 1445 1450

FIG. 7D